

 R^1 represents $C_{1\cdot 12}$ alkyl, $\cdot (CH_2)_a \cdot aryl$, or $(CH_2)_a \cdot Het^1$ (all of which are optionally substituted by one or more substituents selected from ·OH, halo, cyano, nitro, $C_{1\cdot 4}$ alkyl and/or $C_{1\cdot 4}$ alkoxy);

a represents 0, 1, 2, 3, or 4;

Het¹ represents a five to ten-membered heterocyclic ring containing one or more heteroatoms selected from oxygen, nitrogen and/or sulfur, and which also optionally includes one or more =O substituents;

X represents O or S;

 R^{5a} and R^{5b} independently represent H or $C_{1\cdot 3}$ alkyl;

Rf and R³ independently represent H, $C_{1.4}$ alkyl (optionally substituted with one or more nitro or cyano groups), OR^7 , $N(R^{7a})R^{7b}$, $OC(O)R^8$ or together form $-O-(CH_2)_2-O-$, $(CH_2)_3-$, $-(CH_2)_4-$ or $-(CH_2)_5-$;

 R^7 and R^8 independently represent H, $C_{1.6}$ alkyl or $-(CH_2)_b$ -aryl (which latter two groups are optionally substituted by one or more substituents selected from -OH, halo, cyano, nitro $C_{1.4}$ alkyl and/or $C_{1.4}$ alkoxy);

R^{7a} and R^{7b} independently represent H or C₁₋₆ alkyl;

b represents 0, 1, 2, 3 or 4;

R4 represents H or C_{1.6} alkyl;

D represents H, -OH, or $(CH_2)_cN(R^{10})(R^{11})$;

c represents 0, 1, 2, 3 or 4;

 $R^{10} \text{ represents H, C}_{1.6} \text{ alkyl, -(CH}_2)_d\text{-aryl, -C(NH)NH}_2, -S(O)_2 R^{13}, \\ -[C(O)]_e N(R^{14})(R^{15}), -C(O)R^{16} \text{ or -C(O)OR}^{17},$

e represents 1 or 2;

 R^{11} represents H, $C_{1.6}$ alkyl, $-C(O)R^{18}$ or $-(CH_2)_f$ -aryl (which latter group is optionally substituted by one or more substituents selected from -OH, cyano, halo, amino, nitro, $C_{1.6}$ alkyl and/or $C_{1.6}$ alkoxy);

 R^{14} , R^{15} , R^{16} , R^{17} and R^{18} independently represent H, $C_{1.6}$ alkyl, Het² or $\cdot (CH_2)_g$ -aryl (which latter three groups are optionally substituted by one or more substituents selected from -OH, cyano, halo, amino, nitro, $C_{1.6}$ alkyl and/or $C_{1.6}$ alkoxy);

Sih

B/2

 R^{13} represents $C_{1.6}$ alkyl, aryl or $(CH_2)_h$ -aryl (all of which are all optionally substituted by one or more substituents chosen from halo, nitro, $C_{1.6}$ alkyl and/or $C_{1.6}$ alkoxy).

d, f, g and h independently represent 0, 1, 2, 3 or 4;

Het² represents a five to ten-membered heterocyclic ring containing one or more heteroatoms selected from oxygen, nitrogen and/or sulfur, and which also optionally includes one or more =0 substituents;

 R^6 represents one or more optional substituents selected from -OH, cyano, halo, amino, nitro, $C_{1.6}$ alkyl (optionally terminated by -N(H)C(O)OR^{18a}),

 $C_{1.6}$ alkoxy, $\cdot C(O)N(H)R^{19}$, $\cdot NHC(O)N(H)R^{20}$, $\cdot N(H)S(O)_2R^{21}$ and $\cdot OS(O)_2R^{22}$;

 R^{19} and R^{20} independently represent H or $C_{1.6}$ alkyl;

R^{18a}, R²¹ and R²² independently represent C_{1.6} alkyl;

A represents a single bond, C_1 alkylene, $\cdot N(R^{23})(CH_2)_{j^-}$, $\cdot O(CH_2)_{j^-}$ or $\cdot (CH_2)_J C(H)(OR^{23})(CH_2)_{k^-}$ (in which latter three groups, the $\cdot (CH_2)_{j^-}$ group is attached to the bispidine nitrogen atom, and which latter four groups are all optionally substituted by one or more OH groups);

B represents a single bond, $C_{1\cdot4}$ alkylene, $\cdot(CH_2)_mN(R^{24})\cdot$, $(CH_2)_mS(O)_n\cdot$, $\cdot(CH_2)_mO\cdot$ (in which three latter groups, the $\sim(CH_2)_m$ - group is attached to the carbon atom bearing D and R⁴), $\cdot C(O)N(R^{24})\cdot$ (in which latter group, the $\cdot C(O)\cdot$ group is attached to the carbon atom bearing D and R⁴), $N(R^{24})C(O)O(CH_2)_m\cdot$ or $\cdot N(R^{24})(CH_2)_m\cdot$ (in which latter two groups, the $N(R^{24})$ group is attached to the carbon atom bearing D and R⁴);

j, k and m independently represent 0, 1, 2, 3 or 4;

(

n represents 0, 1 or 2;

R³ represents H, C_{1.6} alkyl or

R²⁴ represents H or C_{1.6} alkyl;

 R^{25} represents H, $C_{1.6}$ alkyl, Het^3 or $\cdot(CH_2)_p$ -aryl (which latter two groups are optionally substituted by one or more substituents selected from -OH, cyano, halo, amino, nitro, $C_{1.6}$ alkyl and/or $C_{1.6}$ alkoxy);

Het³ represents a five to ten-membered heterocyclic ring containing one or more heteroatoms-selected from oxygen, nitrogen and/or sulfur, and which also optionally includes one or more =0 substituents;

p represents 0, 1, 2, 3 &r 4;

or a pharmaceutically acceptable salt, solvate or protected derivative thereof;

provided that:

- (a) when D represents either H $^{\circ}$ OH, and R^{5a} and R^{5b} both represent H, then at least one of R² and R³ represents OR OC(O)R⁸ or C_{1.4} alkyl, which alkyl group is substituted with one or more nitro or cyano groups; and
- (b) when D represents -OH or -(CH₂)_cN(R¹⁰)R¹¹ in which c represents 0, then:-
- (i) A does not represent $-N(R^{23})(CH_2)_j$, $-O(CH_2)_j$ or $-CH_2)_JC(H)(OR^{23})(CH_2)_k$ (in which k is 0); and/or
- (ii) m does not represent 0 when B represents $\cdot (CH_2)_m N(R^{24}) \cdot ,$ $\cdot (CH_2)_m S(O)_n \cdot \text{ or } \cdot (CH_2)_m O \cdot .$

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5.b

23 (Amended). A compound of formula IV as defined in Claim 21, or a protected derivative thereof, provided that when R^{5a} and R^{5b} both represent H, then at least one of R^2 and R^3 represents OR^7 , $OC(O)R^8$ or $C_{1.4}$ alkyl, which alkyl group is substituted with one or more nitro or cyano groups.

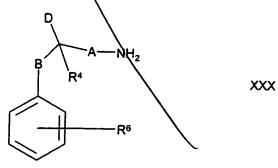
26 (Amended). A process for the preparation of a compound of formula VIII, as defined in Claim 24, or a compound of formula XVII, as defined in Claim 25, which comprises reaction of a compound of formula XXIX,

Sub

XXIX

wherein R^Z represents H or $-C(O)XR^1$ and R^1 , R^{5a} , R^{5b} and X are as defined in Claim 1 with [(as appropriate) either:

(1)] a compound of formula XXX,



or a protected derivative thereof, wherein R⁴, R⁶, A, B and D are as defined in Claim 1, in all cases in the presence of a formaldehyde.

Please add the following new claim.

an atrial or a ventricular arrhythmia.